

CLAIMS

- 1 1. A method of studying the infectivity of a pathogen in tissues comprising the steps of:
 - 2 isolating host cells;
 - 3 placing said isolated host cells into a bioreactor comprising culture medium;
 - 4 applying sedimental shear stress to the cells in the cell culture to form a three-dimensional
 - 5 tissue mass;
 - 6 seeding the formed tissue mass in a tissue culture vessel;
 - 7 introducing an infectious pathogen into said three-dimensional mass; and
 - 8 assaying the infectivity of said infectious pathogen.
- 1 2. The method of claim 1, optionally comprising a culture matrix that facilitates the growth of
- 2 said host cells.
- 1 3. The method of claim 1, wherein the bioreactor is a rotating wall vessel.
- 1 4. The method of claim 1, wherein said isolated host cells are epithelial cells.
- 1 5. The method of claim 4, wherein said epithelial cells are human intestinal cells.
- 1 6. The method of claim 2, wherein said culture matrix consists of microbeads or microcarriers.
- 1 7. The method of claim 1, wherein said infectious pathogen is selected from the group consisting
- 2 essentially of viruses, bacteria, protozoa, parasites and fungi.
- 1 8. The method of claim 7, wherein said infectious pathogen is *Salmonella typhimurium*.
- 1 9. The method of claim 1, wherein said culture medium comprises fetal bovine serum and a tri-
- 2 sugar based medium selected from the mixtures of the group consisting of fructose, galactose and
- 3 lactose.
- 1 10. The method of claim 6, wherein said microbeads are collagen-coated microbeads.

1 11. A method of studying the infectivity of a pathogen in tissues comprising the steps of:
2 isolating intestinal epithelial cells;
3 placing said intestinal epithelial cells into a bioreactor comprising culture medium;
4 applying sedimental shear stress to the cells in the cell culture to form a three-dimensional
5 tissue mass;
6 seeding the formed tissue mass in a tissue culture vessel; and,
7 introducing an infectious pathogen to the formed tissue mass.

1 12. The method of claim 11, wherein said infectious pathogen is *Salmonella typhimurium*.

1 13. A method of measuring the chemosensitivity of tissues to a toxic materials comprising:
2 isolating host cells;
3 placing said isolated host cells into a bioreactor comprising culture medium;
4 applying sedimental shear stress to the cells in the cell culture to form a three-
5 dimensional tissue mass;
6 seeding the formed tissue mass in a tissue culture vessel;
7 introducing a toxic material into said three-dimensional tissue mass; and
8 assaying the chemosensitivity of said toxic material.

1 14. The method of claim 13, optionally comprising a culture matrix that facilitates the growth of
2 said host cells.

1 15. The method of claim 13, wherein said isolated host cells are epithelial cells.

1 16. The method of claim 15, wherein said epithelial cells are human renal cells.

1 17. The method of claim 14, wherein said culture matrix consists of microbeads or
2 microcarriers.

1 18. The method of claim 17, wherein said microbeads are collagen-coated microbeads.

- 1 19. The method of claim 13, wherein said toxic material is a chemotherapeutic material.
- 1 20. The method of claim 19, wherein said chemotherapeutic material is an antibiotic.
- 1 21. The method of claim 20, wherein said antibiotic is gentamicin.
- 1 22. The method of claim 13, wherein said culture medium comprises fetal bovine serum and
2 DMEM/F12.
- 1 23. A method of measuring the chemosensitivity of tissues to a toxic materials comprising:
2 isolating human renal epithelial cells;
3 placing said isolated human renal epithelial cells into a bioreactor comprising culture
4 medium;
5 applying sedimental shear stress to the cells in the cell culture to form a three-
6 dimensional tissue mass;
7 seeding the formed tissue mass in a tissue culture vessel;
8 treating the three dimensional tissue mass with a toxic material; and
9 assaying the chemosensitivity of said toxic material.
- 1 24. The method of claim 23, wherein said toxic material is a chemotherapeutic material.
- 2 25. The method of claim 24, wherein said chemotherapeutic material is an antibiotic.